

Experiment Number: 91005-03

Test Type: 90-DAY

Route: Gavage

Species/Strain: Rat/F 344/N

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Test Compound: Methylene blue trihydrate

CAS Number: 7220-79-3

Date Report Requested: 10/15/2014

Time Report Requested: 18:02:39

First Dose M/F: NA / NA

Lab: BAT

C Number: C91005B

Lock Date: 05/12/1994

Cage Range: All

Date Range: All

Reasons For Removal: All

Removal Date Range: All

Treatment Groups: All

Study Gender: Both

PWG Approval Date NONE

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First Dose M/F: NA / NA

Lab: BAT

**F 344/N Rat Male
0 MG/KG**

DAY ON TEST	0 0 0 0 0 0 0 0 0 0										
	0 0 0 0 0 0 0 0 0 0										
ANIMAL ID	9 9 9 9 9 9 9 9 9 9										
	2 2 2 2 2 2 2 2 2 2										
	0 0 0 0 0 0 0 0 0 0										
	0 0 0 0 0 0 0 0 0 0										
	0 0 0 0 0 0 0 0 0 0										
	0 0 0 0 0 0 0 0 0 0										
	1 2 3 4 5 6 7 8 9 0										
*TOTALS											

Alimentary System

Esophagus	+	+	+	+	+	+	+	+	+	+	10
Intestine Large, Cecum	+	+	+	+	+	+	+	+	+	+	10
Intestine Large, Colon	+	+	+	+	+	+	+	+	+	+	10
Intestine Large, Rectum	+	+	+	+	+	+	+	+	+	+	10
Intestine Small, Duodenum	+	+	+	+	+	+	+	+	+	+	10
Intestine Small, Ileum	+	+	+	+	+	+	+	+	+	+	10
Intestine Small, Jejunum	+	+	+	+	+	+	+	+	+	+	10
Liver	+	+	+	+	+	+	+	+	+	+	10
Centrilobular, Vacuolization Cytoplasmic	2	1	1	2	2	1	1	2		8	1.5
Hepatodiaphragmatic Nodule	X					X				2	
Mesentery						+				1	
Fat, Inflammation, Chronic, Focal						2				1	2.0
Pancreas	+	+	+	+	+	+	+	+	+	+	10
Salivary Glands	+	+	+	+	+	+	+	+	+	+	10
Stomach, Forestomach	+	+	+	+	+	+	+	+	+	+	10
Stomach, Glandular	+	+	+	+	+	+	+	+	+	+	10

Cardiovascular System

Blood Vessel	+	+	+	+	+	+	+	+	+	+	10
Heart	+	+	+	+	+	+	+	+	+	+	10
Inflammation, Chronic, Diffuse				1					1	1.0	
Inflammation, Chronic, Focal	1	1	1	1	1	1			5	1.0	

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ ..Tissue examined microscopically

M ..Missing tissue

X ..Lesion present

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I ..Insufficient tissue

BLANK ..Not examined microscopically

1-4 ..Lesion qualified as:

1) Minimal 3) Moderate

2) Mild 4) Marked

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First Dose M/F: NA / NA

Lab: BAT

**F 344/N Rat Male
0 MG/KG**

DAY ON TEST	0 0 0 0 0 0 0 0 0 0									
	0 0 0 0 0 0 0 0 0 0									
ANIMAL ID	9	9	9	9	9	9	9	9	9	9
	2	2	2	2	2	2	2	2	2	2
	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	1
	1	2	3	4	5	6	7	8	9	0

*TOTALS

Endocrine System

Adrenal Cortex	+	+	+	+	+	+	+	+	+	+	10
Adrenal Medulla	+	+	+	+	+	+	+	+	+	+	10
Islets, Pancreatic	+	+	+	+	+	+	+	+	+	+	10
Parathyroid Gland	+	M	+	+	M	M	+	M	+	+	6
Pituitary Gland	+	+	+	+	+	+	+	+	+	+	10
Thyroid Gland	+	+	+	+	+	+	+	+	+	+	10

General Body System

NONE

Genital System

Epididymis	+	+	+	+	+	+	+	+	+	+	10
Preputial Gland	+	+	+	+	+	+	+	+	+	+	10
Prostate	+	+	+	+	+	+	+	+	+	+	10
Inflammation, Chronic Active	1	1	1								3 1.0
Seminal Vesicle	+	+	+	+	+	+	+	+	+	+	10
Testes	+	+	+	+	+	+	+	+	+	+	10

Hematopoietic System

Bone Marrow	+	+	+	+	+	+	+	+	+	+	10
Lymph Node											1
Mediastinal, Hyperplasia, Lymphoid, Diffuse							3				1 3.0
Lymph Node, Mandibular	+	+	+	+	+	+	+	+	+	+	10

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

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First Dose M/F: NA / NA

Lab: BAT

**F 344/N Rat Male
0 MG/KG**

DAY ON TEST	0 0 0 0 0 0 0 0 0 0									
	0 0 0 0 0 0 0 0 0 0									
ANIMAL ID	9 9 9 9 9 9 9 9 9 9									
	2 2 2 2 2 2 2 2 2 2									
	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	1
	1	2	3	4	5	6	7	8	9	0

***TOTALS**

Lymph Node, Mandibular	+	+	+	+	+	+	+	+	+	+	10
Lymph Node, Mesenteric	+	+	+	+	+	+	+	+	+	+	10
Spleen	+	+	+	+	+	+	+	+	+	+	10
Capsule, Fibrosis							1				1 1.0
Thymus	+	+	+	+	+	+	+	+	+	+	10
Integumentary System											
Mammary Gland	+	+	+	+	+	+	+	+	+	+	10
Skin	+	+	+	+	+	+	+	+	+	+	10
Musculoskeletal System											
Bone	+	+	+	+	+	+	+	+	+	+	10
Nervous System											
Brain	+	+	+	+	+	+	+	+	+	+	10
Respiratory System											
Lung	+	+	+	+	+	+	+	+	+	+	10
Alveolus, Infiltration Cellular, Macrophage						1					1 1.0
Inflammation, Chronic, Diffuse							2	2	2	1	2 1.8
Inflammation, Focal					1	1					2 1.0
Nose	+	+	+	+	+	+	+	+	+	+	10
Trachea	+	+	+	+	+	+	+	+	+	+	10

Special Senses System

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M ..Missing tissue

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Time Report Requested: 18:02:39

First Dose M/F: NA / NA

Lab: BAT

**F 344/N Rat Male
0 MG/KG**

DAY ON TEST										
	0	0	0	0	0	0	0	0	0	0
ANIMAL ID	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	1
	1	2	3	4	5	6	7	8	9	0

*TOTALS

NONE

Urinary System

Kidney	+	+	+	+	+	+	+	+	+	+	10
Nephropathy			1	1			1	1			4 1.0
Urinary Bladder	+	+	+	+	+	+	+	+	+	+	10

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+ ..Tissue examined microscopically

M ..Missing tissue

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Route: Gavage

Species/Strain: Rat/F 344/N

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Test Compound: Methylene blue trihydrate

CAS Number: 7220-79-3

Date Report Requested: 10/15/2014

Time Report Requested: 18:02:39

First Dose M/F: NA / NA

Lab: BAT

F 344/N Rat Male
25 MG/KG

DAY ON TEST										
	0	0	0	0	0	0	0	0	0	0
ANIMAL ID	0	0	0	0	0	0	0	0	0	0
	9	9	9	9	9	9	7	9	9	9
	2	2	2	2	2	2	1	2	2	2
	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0
	1	1	1	1	1	1	1	1	1	2
	1	2	3	4	5	6	7	8	9	0

*TOTALS

Alimentary System

Esophagus	+	1
Intestine Large, Cecum	+	1
Intestine Large, Colon	+	1
Intestine Large, Rectum	+	1
Intestine Small, Duodenum	+	1
Intestine Small, Ileum	+	1
Intestine Small, Jejunum	+	1
Liver	+	2
Centrilobular, Vacuolization Cytoplasmic	1	1 1.0
Hepatodiaphragmatic Nodule	X	1
Pancreas	+	1
Salivary Glands	+	1
Stomach, Forestomach	+	1
Stomach, Glandular	+	1

Cardiovascular System

Blood Vessel	+	1
Heart	+	1

Endocrine System

Adrenal Cortex	+	1
Adrenal Medulla	+	1

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ ..Tissue examined microscopically

M ..Missing tissue

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Test Type: 90-DAY

Route: Gavage

Species/Strain: Rat/F 344/N

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Test Compound: Methylene blue trihydrate

CAS Number: 7220-79-3

Date Report Requested: 10/15/2014

Time Report Requested: 18:02:40

First Dose M/F: NA / NA

Lab: BAT

**F 344/N Rat Male
25 MG/KG**

DAY ON TEST										
	0	0	0	0	0	0	0	0	0	0
ANIMAL ID	0	0	0	0	0	0	0	0	0	0
	9	9	9	9	9	9	7	9	9	9
	2	2	2	2	2	2	1	2	2	2
	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0
	1	1	1	1	1	1	1	1	1	2
	1	2	3	4	5	6	7	8	9	0

*TOTALS

Islets, Pancreatic

+

1

Parathyroid Gland

+

1

Pituitary Gland

+

1

Thyroid Gland

+

1

General Body System

NONE

Genital System

Epididymis

+

1

Preputial Gland

+

1

Prostate

+

1

Inflammation, Chronic Active

1

1 1.0

Seminal Vesicle

+

1

Testes

+

1

Hematopoietic System

Bone Marrow

++ + + + + + + + + +

10

Lymph Node, Mandibular

+

1

Lymph Node, Mesenteric

+

1

Spleen

++ + + + + + + + + +

10

Capsule, Fibrosis

1 1

2 1.0

Congestion

1 1 2 2 1 2

2 1 2

Thymus

+

1

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CAS Number: 7220-79-3

Date Report Requested: 10/15/2014

Time Report Requested: 18:02:40

First Dose M/F: NA / NA

Lab: BAT

**F 344/N Rat Male
25 MG/KG**

DAY ON TEST										
	0	0	0	0	0	0	0	0	0	0
ANIMAL ID	0	0	0	0	0	0	0	0	0	0
	9	9	9	9	9	9	7	9	9	9
	2	2	2	2	2	2	1	2	2	2
	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0
	1	1	1	1	1	1	1	1	1	2
	1	2	3	4	5	6	7	8	9	0

*TOTALS

Integumentary System

Mammary Gland	+	1
Skin	+	1

Musculoskeletal System

Bone	+	1
------	---	---

Nervous System

Brain	+	1
-------	---	---

Respiratory System

Lung	+	1
Nose	+	1
Trachea	+	1

Special Senses System

NONE

Urinary System

Kidney	+	1
Nephropathy	1	1 1.0
Vacuolization Cytoplasmic, Diffuse	2	1 2.0
Urinary Bladder	+	1

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Species/Strain: Rat/F 344/N

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Test Compound: Methylene blue trihydrate

CAS Number: 7220-79-3

Date Report Requested: 10/15/2014

Time Report Requested: 18:02:40

First Dose M/F: NA / NA

Lab: BAT

**F 344/N Rat Male
50 MG/KG**

DAY ON TEST										
	0	0	0	0	0	0	0	0	0	0
ANIMAL ID	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0
	2	2	2	2	2	2	2	2	2	2
	1	2	3	4	5	6	7	8	9	0

*TOTALS

Alimentary System

NONE

Cardiovascular System

NONE

Endocrine System

NONE

General Body System

NONE

Genital System

NONE

Hematopoietic System

Bone Marrow

+ + + + + + + + + + 10

Hyperplasia

2 2 2 2 2 2 2 2 2 2 8 2.0

Spleen

+ + + + + + + + + + 10

Congestion

2 2 2 2 2 2 2 2 2 2 10 2.0

Hematopoietic Cell Proliferation

2 2 2 2 2 2 2 2 2 2 10 2.0

Integumentary System

NONE

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+ ..Tissue examined microscopically

M ..Missing tissue

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I ..Insufficient tissue

BLANK ..Not examined microscopically

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Time Report Requested: 18:02:40

First Dose M/F: NA / NA

Lab: BAT

**F 344/N Rat Male
50 MG/KG**

| DAY ON TEST | | | | | | | | | | |
|-------------|---|---|---|---|---|---|---|---|---|---|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| ANIMAL ID | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 |

*TOTALS

Musculoskeletal System

NONE

Nervous System

NONE

Respiratory System

Lung

+ 1

Cyst

3 1 3.0

Special Senses System

NONE

Urinary System

NONE

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Date Report Requested: 10/15/2014

Time Report Requested: 18:02:40

First Dose M/F: NA / NA

Lab: BAT

**F 344/N Rat Male
100 MG/KG**

| DAY ON TEST | | | | | | | | | | |
|-------------|---|---|---|---|---|---|---|---|---|---|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| ANIMAL ID | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 4 |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 |

*TOTALS

Alimentary System

Liver

+ +

2

Hepatodiaphragmatic Nodule

X X

2

Cardiovascular System

NONE

Endocrine System

NONE

General Body System

NONE

Genital System

NONE

Hematopoietic System

Bone Marrow

++ + + + + + + + +

10

Hyperplasia

2 2 2 2 2 2 2 2 2 2

10 2.0

Spleen

++ + + + + + + + +

10

Capsule, Fibrosis

1

1 1.0

Congestion

2 2 2 2 2 2 2 2 2 2

10 2.0

Hematopoietic Cell Proliferation

2 2 2 2 2 2 2 2 2 2

9 2.0

Lymph Follic, Depletion Cellular

1 1

2 1.0

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First Dose M/F: NA / NA

Lab: BAT

**F 344/N Rat Male
100 MG/KG**

| DAY ON TEST | | | | | | | | | | |
|-------------|---|---|---|---|---|---|---|---|---|---|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| ANIMAL ID | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 4 |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 |

*TOTALS

Integumentary System

NONE

Musculoskeletal System

NONE

Nervous System

NONE

Respiratory System

NONE

Special Senses System

Eye

+

1

Urinary System

NONE

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+ ..Tissue examined microscopically

M ..Missing tissue

X ..Lesion present

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I ..Insufficient tissue

BLANK ..Not examined microscopically

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CAS Number: 7220-79-3

Date Report Requested: 10/15/2014

Time Report Requested: 18:02:40

First Dose M/F: NA / NA

Lab: BAT

**F 344/N Rat Male
200 MG/KG**

| DAY ON TEST | | | | | | | | | | |
|-------------|---|---|---|---|---|---|---|---|---|---|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| ANIMAL ID | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 5 |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 |

TOTALS*Alimentary System**

| | | | | | | | | | | | |
|--|---|---|---|---|---|---|---|---|---|---|--------|
| Esophagus | + | + | + | + | + | + | + | + | + | + | 10 |
| Intestine Large, Cecum | + | + | + | + | + | + | + | + | + | + | 10 |
| Intestine Large, Colon | + | + | + | + | + | + | + | + | + | + | 10 |
| Intestine Large, Rectum | + | + | + | + | + | + | + | + | + | + | 10 |
| Intestine Small, Duodenum | + | + | + | + | + | + | + | + | + | + | 10 |
| Intestine Small, Ileum | + | + | + | + | + | + | + | + | + | + | 10 |
| Intestine Small, Jejunum | + | + | + | + | + | + | + | + | + | + | 10 |
| Liver | + | + | + | + | + | + | + | + | + | + | 10 |
| Centrilobular, Vacuolization Cytoplasmic | 2 | 2 | 2 | 2 | 1 | 1 | 1 | 1 | 1 | 1 | 10 1.4 |
| Pancreas | + | + | + | + | + | + | + | + | + | + | 10 |
| Salivary Glands | + | + | + | + | + | + | + | + | + | + | 10 |
| Stomach, Forestomach | + | + | + | + | + | + | + | + | + | + | 10 |
| Stomach, Glandular | + | + | + | + | + | + | + | + | + | + | 10 |
| Mineralization | | | | | 1 | | | | | | 1 1.0 |

Cardiovascular System

| | | | | | | | | | | | |
|--------------------------------|---|---|---|---|---|---|---|---|---|---|-------|
| Blood Vessel | + | + | + | + | + | + | + | + | + | + | 10 |
| Heart | + | + | + | + | + | + | + | + | + | + | 10 |
| Inflammation, Chronic, Diffuse | 1 | | | | 2 | 1 | | | | | 3 1.3 |
| Inflammation, Chronic, Focal | | | | 1 | 1 | | 1 | | | | 3 1.0 |

Endocrine System

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ ..Tissue examined microscopically

M ..Missing tissue

X ..Lesion present

A ..Autolysis precludes evaluation

I ..Insufficient tissue

BLANK ..Not examined microscopically

1-4 ..Lesion qualified as:

1) Minimal 3) Moderate

2) Mild 4) Marked

Experiment Number: 91005-03

Test Type: 90-DAY

Route: Gavage

Species/Strain: Rat/F 344/N

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Test Compound: Methylene blue trihydrate

CAS Number: 7220-79-3

Date Report Requested: 10/15/2014

Time Report Requested: 18:02:40

First Dose M/F: NA / NA

Lab: BAT

F 344/N Rat Male
200 MG/KG

| DAY ON TEST | | | | | | | | | | |
|-------------|---|---|---|---|---|---|---|---|---|---|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| ANIMAL ID | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 5 |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 |

***TOTALS**

| | | | | | | | | | | | |
|----------------------------------|---|---|---|---|---|---|---|---|---|---|-----|
| Adrenal Cortex | + | + | + | + | + | + | + | + | + | + | 10 |
| Vacuolization Cytoplasmic, Focal | | | | | | | 1 | | | 1 | 1.0 |
| Adrenal Medulla | + | + | + | + | + | + | + | + | + | + | 10 |
| Islets, Pancreatic | + | + | + | + | + | + | + | + | + | + | 10 |
| Parathyroid Gland | + | M | M | M | + | + | + | + | + | + | 7 |
| Pituitary Gland | + | + | + | + | + | + | + | + | + | + | 10 |
| Thyroid Gland | + | + | + | + | + | + | + | + | + | + | 10 |

General Body System

NONE

Genital System

| | | | | | | | | | | | |
|------------------------------|---|---|---|---|---|---|---|---|---|---|-----|
| Epididymis | + | + | + | + | + | + | + | + | + | + | 10 |
| Preputial Gland | + | + | + | + | + | + | + | + | + | + | 10 |
| Prostate | + | + | + | + | + | + | + | + | + | + | 10 |
| Inflammation, Chronic Active | | | | 1 | | 1 | 1 | | | 3 | 1.0 |
| Seminal Vesicle | + | + | + | + | + | + | + | + | + | + | 10 |
| Testes | + | + | + | + | + | + | + | + | + | + | 10 |

Hematopoietic System

| | | | | | | | | | | | |
|------------------------|---|---|---|---|---|---|---|---|---|---|----|
| Bone Marrow | + | + | + | + | + | + | + | + | + | + | 10 |
| Hyperplasia | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 10 |
| Lymph Node, Mandibular | + | + | + | + | + | + | + | + | + | + | 10 |
| Lymph Node, Mesenteric | + | + | + | + | + | + | + | + | + | + | 10 |

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ ..Tissue examined microscopically

M ..Missing tissue

X ..Lesion present

A ..Autolysis precludes evaluation

I ..Insufficient tissue

BLANK ..Not examined microscopically

1-4 ..Lesion qualified as:

- 1) Minimal 3) Moderate
- 2) Mild 4) Marked

Experiment Number: 91005-03

Test Type: 90-DAY

Route: Gavage

Species/Strain: Rat/F 344/N

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Test Compound: Methylene blue trihydrate

CAS Number: 7220-79-3

Date Report Requested: 10/15/2014

Time Report Requested: 18:02:40

First Dose M/F: NA / NA

Lab: BAT

**F 344/N Rat Male
200 MG/KG**

| DAY ON TEST | 0 0 0 0 0 0 0 0 0 0 | | | | | | | | | |
|-------------|---------------------|--|--|--|--|--|--|--|--|--|
| | 0 0 0 0 0 0 0 0 0 0 | | | | | | | | | |
| ANIMAL ID | 9 9 9 9 9 9 9 9 9 9 | | | | | | | | | |
| | 2 2 2 2 2 2 2 2 2 2 | | | | | | | | | |
| | 0 0 0 0 0 0 0 0 0 0 | | | | | | | | | |
| | 0 0 0 0 0 0 0 0 0 0 | | | | | | | | | |
| | 0 0 0 0 0 0 0 0 0 0 | | | | | | | | | |
| | 4 4 4 4 4 4 4 4 4 5 | | | | | | | | | |
| | 1 2 3 4 5 6 7 8 9 0 | | | | | | | | | |

***TOTALS**

| | | | | | | | | | | | |
|--------|---|---|---|---|---|---|---|---|---|---|----|
| Spleen | + | + | + | + | + | + | + | + | + | + | 10 |
|--------|---|---|---|---|---|---|---|---|---|---|----|

| | | | | | | | | | | | |
|-------------------|---|---|---|---|---|---|---|---|---|---|-------|
| Capsule, Fibrosis | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 9 1.0 |
|-------------------|---|---|---|---|---|---|---|---|---|---|-------|

| | | | | | | | | | | | |
|------------|---|---|---|---|---|---|---|---|---|---|--------|
| Congestion | 2 | 2 | 2 | 1 | 2 | 2 | 2 | 2 | 2 | 2 | 10 1.9 |
|------------|---|---|---|---|---|---|---|---|---|---|--------|

| | | | | | | | | | | | |
|----------------------------------|---|---|---|---|---|---|---|---|---|---|--------|
| Hematopoietic Cell Proliferation | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 10 2.0 |
|----------------------------------|---|---|---|---|---|---|---|---|---|---|--------|

| | | | | | | | | | | | |
|----------------------------------|---|---|---|---|---|---|--|--|--|--|-------|
| Lymph Follic, Depletion Cellular | 2 | 2 | 1 | 1 | 1 | 1 | | | | | 5 1.4 |
|----------------------------------|---|---|---|---|---|---|--|--|--|--|-------|

| | | | | | | | | | | | |
|--------|---|---|---|---|---|---|---|---|---|---|----|
| Thymus | + | + | + | + | + | + | + | + | + | + | 10 |
|--------|---|---|---|---|---|---|---|---|---|---|----|

Integumentary System

| | | | | | | | | | | | |
|---------------|---|---|---|---|---|---|---|---|---|---|----|
| Mammary Gland | + | + | + | + | + | + | + | + | + | + | 10 |
|---------------|---|---|---|---|---|---|---|---|---|---|----|

| | | | | | | | | | | | |
|------|---|---|---|---|---|---|---|---|---|---|----|
| Skin | + | + | + | + | + | + | + | + | + | + | 10 |
|------|---|---|---|---|---|---|---|---|---|---|----|

Musculoskeletal System

| | | | | | | | | | | | |
|------|---|---|---|---|---|---|---|---|---|---|----|
| Bone | + | + | + | + | + | + | + | + | + | + | 10 |
|------|---|---|---|---|---|---|---|---|---|---|----|

Nervous System

| | | | | | | | | | | | |
|-------|---|---|---|---|---|---|---|---|---|---|----|
| Brain | + | + | + | + | + | + | + | + | + | + | 10 |
|-------|---|---|---|---|---|---|---|---|---|---|----|

Respiratory System

| | | | | | | | | | | | |
|------|---|---|---|---|---|---|---|---|---|---|----|
| Lung | + | + | + | + | + | + | + | + | + | + | 10 |
|------|---|---|---|---|---|---|---|---|---|---|----|

| | | | | | | | | | | | |
|--------------------------------|--|--|--|--|--|--|---|--|--|--|-------|
| Inflammation, Chronic, Diffuse | | | | | | | 2 | | | | 1 2.0 |
|--------------------------------|--|--|--|--|--|--|---|--|--|--|-------|

| | | | | | | | | | | | |
|------|---|---|---|---|---|---|---|---|---|---|----|
| Nose | + | + | + | + | + | + | + | + | + | + | 10 |
|------|---|---|---|---|---|---|---|---|---|---|----|

| | | | | | | | | | | | |
|---------|---|---|---|---|---|---|---|---|---|---|----|
| Trachea | + | + | + | + | + | + | + | + | + | + | 10 |
|---------|---|---|---|---|---|---|---|---|---|---|----|

Special Senses System

NONE

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ ..Tissue examined microscopically

M ..Missing tissue

X ..Lesion present

A ..Autolysis precludes evaluation

I ..Insufficient tissue

BLANK ..Not examined microscopically

1-4 ..Lesion qualified as:

1) Minimal 3) Moderate

2) Mild 4) Marked

Experiment Number: 91005-03

Test Type: 90-DAY

Route: Gavage

Species/Strain: Rat/F 344/N

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Test Compound: Methylene blue trihydrate

CAS Number: 7220-79-3

Date Report Requested: 10/15/2014

Time Report Requested: 18:02:40

First Dose M/F: NA / NA

Lab: BAT

F 344/N Rat Male
200 MG/KG

| DAY ON TEST | | | | | | | | | | |
|-------------|---|---|---|---|---|---|---|---|---|---|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 |
| 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| ANIMAL ID | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 5 |
| 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | |

*TOTALS

Urinary System

| | | | | | | | | | | | |
|-----------------|---|---|---|---|---|---|---|---|---|---|-------|
| Kidney | + | + | + | + | + | + | + | + | + | + | 10 |
| Nephropathy | | | | | 1 | | 1 | | 1 | | 3 1.0 |
| Urinary Bladder | + | + | + | + | + | + | + | + | + | + | 10 |

END OF MALE DATA

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ ..Tissue examined microscopically

M ..Missing tissue

X ..Lesion present

A ..Autolysis precludes evaluation

I ..Insufficient tissue

BLANK ..Not examined microscopically

1-4 ..Lesion qualified as:

1) Minimal 3) Moderate

2) Mild 4) Marked

Experiment Number: 91005-03

Test Type: 90-DAY

Route: Gavage

Species/Strain: Rat/F 344/N

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Test Compound: Methylene blue trihydrate

CAS Number: 7220-79-3

Date Report Requested: 10/15/2014

Time Report Requested: 18:02:40

First Dose M/F: NA / NA

Lab: BAT

**F 344/N Rat Female
0 MG/KG**

| DAY ON TEST | | | | | | | | | | |
|-------------|---|---|---|---|---|---|---|---|---|---|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| ANIMAL ID | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 6 |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 |

TOTALS*Alimentary System**

| | | | | | | | | | | | |
|------------------------------|---|---|---|---|---|---|---|---|---|---|-------|
| Esophagus | + | + | + | + | + | + | + | + | + | + | 10 |
| Intestine Large, Cecum | + | + | + | + | + | + | + | + | + | + | 10 |
| Intestine Large, Colon | + | + | + | + | + | + | + | + | + | + | 10 |
| Intestine Large, Rectum | + | + | + | + | + | + | + | + | + | + | 10 |
| Intestine Small, Duodenum | + | + | + | + | + | + | + | + | + | + | 10 |
| Intestine Small, Ileum | + | + | + | + | + | + | + | + | + | + | 10 |
| Intestine Small, Jejunum | + | + | + | + | + | + | + | + | + | + | 10 |
| Liver | + | + | + | + | + | + | + | + | + | + | 10 |
| Mesentery | + | | | | | | | | | | 1 |
| Inflammation, Chronic, Focal | | | | | | | | | | | 1 2.0 |
| Pancreas | + | + | + | + | + | + | + | + | + | + | 10 |
| Salivary Glands | + | + | + | + | + | + | + | + | + | + | 10 |
| Stomach, Forestomach | + | + | + | + | + | + | + | + | + | + | 10 |
| Stomach, Glandular | + | + | + | + | + | + | + | + | + | + | 10 |

Cardiovascular System

| | | | | | | | | | | | |
|--------------|---|---|---|---|---|---|---|---|---|---|----|
| Blood Vessel | + | + | + | + | + | + | + | + | + | + | 10 |
| Heart | + | + | + | + | + | + | + | + | + | + | 10 |

Endocrine System

| | | | | | | | | | | | |
|-----------------|---|---|---|---|---|---|---|---|---|---|----|
| Adrenal Cortex | + | + | + | + | + | + | + | + | + | + | 10 |
| Adrenal Medulla | + | + | + | + | + | + | + | + | + | + | 10 |

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ ..Tissue examined microscopically

M ..Missing tissue

X ..Lesion present

A ..Autolysis precludes evaluation

I ..Insufficient tissue

BLANK ..Not examined microscopically

1-4 ..Lesion qualified as:

- 1) Minimal 3) Moderate
 2) Mild 4) Marked

Experiment Number: 91005-03

Test Type: 90-DAY

Route: Gavage

Species/Strain: Rat/F 344/N

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Test Compound: Methylene blue trihydrate

CAS Number: 7220-79-3

Date Report Requested: 10/15/2014

Time Report Requested: 18:02:40

First Dose M/F: NA / NA

Lab: BAT

F 344/N Rat Female
0 MG/KG

| DAY ON TEST | | | | | | | | | | |
|-------------|---|---|---|---|---|---|---|---|---|---|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 |
| 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| ANIMAL ID | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 6 |
| 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | |

*TOTALS

| | | | | | | | | | | | |
|--------------------|---|---|---|---|---|---|---|---|---|---|----|
| Islets, Pancreatic | + | + | + | + | + | + | + | + | + | + | 10 |
| Parathyroid Gland | + | + | + | + | + | + | + | + | + | + | 10 |
| Pituitary Gland | + | + | + | + | + | + | + | + | + | + | 10 |
| Thyroid Gland | + | + | + | + | + | + | + | + | + | + | 10 |

General Body System

NONE

Genital System

| | | | | | | | | | | | |
|----------------------|---|---|---|---|---|---|---|---|---|-----|----|
| Clitoral Gland | + | + | + | + | + | + | + | + | + | + | 10 |
| Ovary | + | + | + | + | + | + | + | + | + | + | 10 |
| Periovary Tiss, Cyst | | | | 2 | 3 | | | | 2 | 2.5 | |
| Uterus | + | + | + | + | + | + | + | + | + | + | 10 |
| Dilatation | | | | | | | 1 | 1 | 2 | 1.0 | |

Hematopoietic System

| | | | | | | | | | | | |
|------------------------|---|---|---|---|---|---|---|---|---|---|----|
| Bone Marrow | + | + | + | + | + | + | + | + | + | + | 10 |
| Lymph Node, Mandibular | + | + | + | + | + | + | + | + | + | + | 10 |
| Lymph Node, Mesenteric | + | + | + | + | + | + | + | + | + | + | 10 |
| Spleen | + | + | + | + | + | + | + | + | + | + | 10 |
| Thymus | + | + | + | + | + | + | + | + | + | + | 10 |

Integumentary System

| | | | | | | | | | | | |
|---------------|---|---|---|---|---|---|---|---|---|---|----|
| Mammary Gland | + | + | + | + | + | + | + | + | + | + | 10 |
| Skin | + | + | + | + | + | + | + | + | + | + | 10 |

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ ..Tissue examined microscopically

M ..Missing tissue

X ..Lesion present

A ..Autolysis precludes evaluation

I ..Insufficient tissue

BLANK ..Not examined microscopically

1-4 ..Lesion qualified as:

1) Minimal 3) Moderate

2) Mild 4) Marked

Experiment Number: 91005-03
Test Type: 90-DAY
Route: Gavage
Species/Strain: Rat/F 344/N

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL
Test Compound: Methylene blue trihydrate
CAS Number: 7220-79-3

Date Report Requested: 10/15/2014
Time Report Requested: 18:02:40
First Dose M/F: NA / NA
Lab: BAT

F 344/N Rat Female
0 MG/KG

| | DAY ON TEST | | | | | | | | | | | *TOTALS |
|-----------|-------------|---|---|---|---|---|---|---|---|---|---|----------------|
| | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| ANIMAL ID | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 6 | |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | | |
| | | | | | | | | | | | | |
| | | | | | | | | | | | | |
| | | | | | | | | | | | | |
| | | | | | | | | | | | | |
| | | | | | | | | | | | | |

Skin

+ + + + + + + + + + 10

Musculoskeletal System

Bone

+ + + + + + + + + + 10

Nervous System

Brain

+ + + + + + + + + + 10

Respiratory System

Lung

+ + + + + + + + + + 10

Inflammation, Chronic, Diffuse

2 1 2.0

Nose

+ + + + + + + + + + 10

Trachea

+ + + + + + + + + + 10

Special Senses System

NONE

Urinary System

Kidney

+ + + + + + + + + + 10

Urinary Bladder

+ + + + + + + + + + 10

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ ..Tissue examined microscopically

M ..Missing tissue

X ..Lesion present

A ..Autolysis precludes evaluation

I ..Insufficient tissue

BLANK ..Not examined microscopically

1-4 ..Lesion qualified as:

- 1) Minimal 3) Moderate
- 2) Mild 4) Marked

Experiment Number: 91005-03

Test Type: 90-DAY

Route: Gavage

Species/Strain: Rat/F 344/N

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Test Compound: Methylene blue trihydrate

CAS Number: 7220-79-3

Date Report Requested: 10/15/2014

Time Report Requested: 18:02:40

First Dose M/F: NA / NA

Lab: BAT

**F 344/N Rat Female
25 MG/KG**

| DAY ON TEST | | | | | | | | | | |
|-------------|---|---|---|---|---|---|---|---|---|---|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| ANIMAL ID | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 |
| | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 |

*TOTALS

Alimentary System

NONE

Cardiovascular System

NONE

Endocrine System

NONE

General Body System

NONE

Genital System

NONE

Hematopoietic System

Bone Marrow

 + + + + + + + + + + 10

Spleen

 + + + + + + + + + + 10

Congestion

 2 2 1 2 2 2 2 1 1 9 1.7

Integumentary System

NONE

Musculoskeletal System

NONE

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ ..Tissue examined microscopically

M ..Missing tissue

X ..Lesion present

A ..Autolysis precludes evaluation

I ..Insufficient tissue

BLANK ..Not examined microscopically

1-4 ..Lesion qualified as:

- 1) Minimal 3) Moderate
- 2) Mild 4) Marked

Experiment Number: 91005-03

Test Type: 90-DAY

Route: Gavage

Species/Strain: Rat/F 344/N

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Test Compound: Methylene blue trihydrate

CAS Number: 7220-79-3

Date Report Requested: 10/15/2014

Time Report Requested: 18:02:40

First Dose M/F: NA / NA

Lab: BAT

**F 344/N Rat Female
25 MG/KG**

| DAY ON TEST | | | | | | | | | | |
|-------------|---|---|---|---|---|---|---|---|---|---|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| ANIMAL ID | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 7 |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 |

*TOTALS

Nervous System

NONE

Respiratory System

NONE

Special Senses System

NONE

Urinary System

NONE

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ ..Tissue examined microscopically

M ..Missing tissue

X ..Lesion present

A ..Autolysis precludes evaluation

I ..Insufficient tissue

BLANK ..Not examined microscopically

1-4 ..Lesion qualified as:

- 1) Minimal 3) Moderate
- 2) Mild 4) Marked

Experiment Number: 91005-03
Test Type: 90-DAY
Route: Gavage
Species/Strain: Rat/F 344/N

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL
Test Compound: Methylene blue trihydrate
CAS Number: 7220-79-3

Date Report Requested: 10/15/2014
Time Report Requested: 18:02:40
First Dose M/F: NA / NA
Lab: BAT

**F 344/N Rat Female
50 MG/KG**

| DAY ON TEST | | | | | | | | | | |
|-------------|---|---|---|---|---|---|---|---|---|---|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| ANIMAL ID | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 |
| | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 8 |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 |

***TOTALS**

Alimentary System

NONE

Cardiovascular System

NONE

Endocrine System

NONE

General Body System

NONE

Genital System

Ovary

| | |
|---|-------|
| + | 1 |
| 2 | 1 2.0 |

Hematopoietic System

Bone Marrow

| | |
|---|----|
| + | 10 |
|---|----|

Hyperplasia

| | |
|---|-------|
| 1 | 4 1.3 |
|---|-------|

Spleen

| | |
|---|----|
| + | 10 |
|---|----|

Congestion

| | |
|---|--------|
| 2 | 10 2.0 |
|---|--------|

Hematopoietic Cell Proliferation

| | |
|---|-------|
| 2 | 8 1.9 |
|---|-------|

Integumentary System

NONE

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ ..Tissue examined microscopically

M ..Missing tissue

X ..Lesion present

A ..Autolysis precludes evaluation

I ..Insufficient tissue

BLANK ..Not examined microscopically

1-4 ..Lesion qualified as:

- 1) Minimal 3) Moderate
- 2) Mild 4) Marked

Experiment Number: 91005-03

Test Type: 90-DAY

Route: Gavage

Species/Strain: Rat/F 344/N

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Test Compound: Methylene blue trihydrate

CAS Number: 7220-79-3

Date Report Requested: 10/15/2014

Time Report Requested: 18:02:41

First Dose M/F: NA / NA

Lab: BAT

**F 344/N Rat Female
50 MG/KG**

| DAY ON TEST | | | | | | | | | | |
|-------------|---|---|---|---|---|---|---|---|---|---|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| ANIMAL ID | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 |
| | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 8 |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 |

*TOTALS

Musculoskeletal System

NONE

Nervous System

NONE

Respiratory System

NONE

Special Senses System

NONE

Urinary System

NONE

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ ..Tissue examined microscopically

M ..Missing tissue

X ..Lesion present

A ..Autolysis precludes evaluation

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Test Compound: Methylene blue trihydrate

CAS Number: 7220-79-3

Date Report Requested: 10/15/2014

Time Report Requested: 18:02:41

First Dose M/F: NA / NA

Lab: BAT

F 344/N Rat Female
100 MG/KG

| DAY ON TEST | | | | | | | | | | |
|-------------|---|---|---|---|---|---|---|---|---|---|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 9 | 9 | 1 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 |
| 2 | 2 | 4 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| ANIMAL ID | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 9 |
| 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | * |

*TOTALS

Alimentary System

| | | | |
|----------------------------|---|---|---|
| Esophagus | + | | 1 |
| Intestine Large, Cecum | + | | 1 |
| Intestine Large, Colon | + | | 1 |
| Intestine Large, Rectum | + | | 1 |
| Intestine Small, Duodenum | + | | 1 |
| Intestine Small, Ileum | + | | 1 |
| Intestine Small, Jejunum | + | | 1 |
| Liver | + | + | 2 |
| Hepatodiaphragmatic Nodule | | X | 1 |
| Pancreas | + | | 1 |
| Salivary Glands | + | | 1 |
| Stomach, Forestomach | + | | 1 |
| Stomach, Glandular | + | | 1 |

Cardiovascular System

| | | |
|--------------|---|---|
| Blood Vessel | + | 1 |
| Heart | + | 1 |

Endocrine System

| | | |
|--------------------|---|---|
| Adrenal Cortex | + | 1 |
| Adrenal Medulla | + | 1 |
| Islets, Pancreatic | + | 1 |

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ ..Tissue examined microscopically

M ..Missing tissue

X ..Lesion present

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Test Compound: Methylene blue trihydrate

CAS Number: 7220-79-3

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Time Report Requested: 18:02:41

First Dose M/F: NA / NA

Lab: BAT

**F 344/N Rat Female
100 MG/KG**

| DAY ON TEST | | | | | | | | | | |
|-------------|---|---|---|---|---|---|---|---|---|---|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| ANIMAL ID | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 9 |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 |

***TOTALS**

Parathyroid Gland

+

1

Pituitary Gland

+

1

Thyroid Gland

+

1

General Body System

NONE

Genital System

Ovary

+

1

Uterus

+

1

Hematopoietic System

Bone Marrow

+ + + + + + + + + +

10

9 1.8

1 2 2 1 2 2 2 2 2 2

1

Hyperplasia

+ M

0

Lymph Node, Mandibular

M

0

Lymph Node, Mesenteric

+ + + + + + + + + +

10

Spleen

1 1 1

3 1.0

Capsule, Fibrosis

2 2 2 2 2 2 2 2 2 2

9 2.0

Congestion

2 2 2 2 2 2 2 2 2 2

9 2.0

Hematopoietic Cell Proliferation

2 2 2 2 2 2 2 2 2 2

9 2.0

Lymph Follic, Depletion Cellular

2

1 2.0

Thymus

+

1

Integumentary System

Mammary Gland

+

1

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ ..Tissue examined microscopically

M ..Missing tissue

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Species/Strain: Rat/F 344/N

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Test Compound: Methylene blue trihydrate

CAS Number: 7220-79-3

Date Report Requested: 10/15/2014

Time Report Requested: 18:02:41

First Dose M/F: NA / NA

Lab: BAT

**F 344/N Rat Female
100 MG/KG**

| DAY ON TEST | | | | | | | | | | |
|-------------|---|---|---|---|---|---|---|---|---|---|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| ANIMAL ID | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 9 |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 |

*TOTALS

Mammary Gland

+

1

Skin

+

1

Musculoskeletal System

Bone

+

1

Nervous System

Brain

+

1

Respiratory System

Lung

+

1

Nose

+

1

Trachea

+

1

Special Senses System

NONE

Urinary System

Kidney

+

1

Urinary Bladder

+

1

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ ..Tissue examined microscopically

M ..Missing tissue

X ..Lesion present

A ..Autolysis precludes evaluation

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Experiment Number: 91005-03

Test Type: 90-DAY

Route: Gavage

Species/Strain: Rat/F 344/N

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Test Compound: Methylene blue trihydrate

CAS Number: 7220-79-3

Date Report Requested: 10/15/2014

Time Report Requested: 18:02:41

First Dose M/F: NA / NA

Lab: BAT

**F 344/N Rat Female
200 MG/KG**

| DAY ON TEST | | | | | | | | | | |
|-------------|---|---|---|---|---|---|---|---|---|---|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| ANIMAL ID | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 |
| | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 0 |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 |

TOTALS*Alimentary System**

| | | | | | | | | | | |
|--|---|---|---|---|---|---|---|---|---|-----|
| Esophagus | + | + | + | + | + | + | + | + | + | 10 |
| Intestine Large, Cecum | + | + | + | + | + | + | + | + | + | 10 |
| Intestine Large, Colon | + | + | + | + | + | + | + | + | + | 10 |
| Intestine Large, Rectum | + | + | + | + | + | + | + | + | + | 10 |
| Intestine Small, Duodenum | + | + | + | + | + | + | + | + | + | 10 |
| Intestine Small, Ileum | + | + | + | M | + | + | + | + | + | 9 |
| Intestine Small, Jejunum | + | + | + | + | + | + | + | + | + | 10 |
| Liver | + | + | + | + | + | + | + | + | + | 10 |
| Centrilobular, Vacuolization Cytoplasmic | 1 | 1 | | | | | | | 2 | 1.0 |
| Hepatodiaphragmatic Nodule | | | | | | | X | | | 1 |
| Pancreas | + | + | + | + | + | + | + | + | + | 10 |
| Salivary Glands | + | + | + | + | + | + | + | + | + | 10 |
| Stomach, Forestomach | + | + | + | + | + | + | + | + | + | 10 |
| Stomach, Glandular | + | + | + | + | + | + | + | + | + | 10 |

Cardiovascular System

| | | | | | | | | | | |
|------------------------------|---|---|---|---|---|---|---|---|---|-----|
| Blood Vessel | + | + | + | + | + | + | + | + | + | 10 |
| Heart | + | + | + | + | + | + | + | + | + | 10 |
| Inflammation, Chronic, Focal | | 1 | | | | | | | 1 | 1.0 |

Endocrine System

| | | | | | | | | | | |
|----------------|---|---|---|---|---|---|---|---|---|----|
| Adrenal Cortex | + | + | + | + | + | + | + | + | + | 10 |
|----------------|---|---|---|---|---|---|---|---|---|----|

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M ..Missing tissue

X ..Lesion present

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Experiment Number: 91005-03

Test Type: 90-DAY

Route: Gavage

Species/Strain: Rat/F 344/N

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Test Compound: Methylene blue trihydrate

CAS Number: 7220-79-3

Date Report Requested: 10/15/2014

Time Report Requested: 18:02:41

First Dose M/F: NA / NA

Lab: BAT

**F 344/N Rat Female
200 MG/KG**

| DAY ON TEST | | | | | | | | | | |
|-------------|---|---|---|---|---|---|---|---|---|---|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| ANIMAL ID | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 1 | 9 | 2 | 9 | 9 | 7 | 2 | 9 | 9 | 9 | 9 |
| 4 | 2 | 7 | 2 | 2 | 1 | 7 | 2 | 2 | 2 | 2 |

***TOTALS**

| | | | | | | | | | | | |
|--------------------|---|---|---|---|---|---|---|---|---|---|----|
| Adrenal Medulla | + | + | + | + | + | + | + | + | + | + | 10 |
| Islets, Pancreatic | + | + | + | + | + | + | + | + | + | + | 10 |
| Parathyroid Gland | + | M | + | M | M | + | + | M | + | + | 6 |
| Pituitary Gland | + | + | + | + | + | + | + | + | + | + | 10 |
| Thyroid Gland | + | + | + | + | + | + | + | + | + | + | 10 |

General Body System

NONE

Genital System

| | | | | | | | | | | | |
|----------------|---|---|---|---|---|---|---|---|---|---|-------|
| Clitoral Gland | + | + | + | + | + | + | + | + | + | + | 10 |
| Ovary | + | + | + | + | + | + | + | + | + | + | 10 |
| Uterus | + | + | + | + | + | + | + | + | + | + | 10 |
| Dilatation | | | | | 1 | | | | | | 1 1.0 |

Hematopoietic System

| | | | | | | | | | | | |
|----------------------------------|---|---|---|---|---|---|---|---|---|---|--------|
| Bone Marrow | + | + | + | + | + | + | + | + | + | + | 10 |
| Hyperplasia | 3 | 3 | 3 | 3 | 3 | 3 | 2 | 3 | 3 | 3 | 10 2.9 |
| Lymph Node, Mandibular | + | + | + | + | + | + | + | + | M | + | 9 |
| Lymph Node, Mesenteric | + | + | + | + | + | + | + | + | + | + | 10 |
| Spleen | + | + | + | + | + | + | + | + | + | + | 10 |
| Capsule, Fibrosis | 1 | 1 | 1 | 1 | | | 1 | 1 | 1 | | 7 1.0 |
| Congestion | 2 | 2 | 1 | 2 | 2 | | 1 | 2 | 2 | 2 | 9 1.8 |
| Hematopoietic Cell Proliferation | 3 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 1 | 2 | 10 2.0 |

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ ..Tissue examined microscopically

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X ..Lesion present

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Experiment Number: 91005-03

Test Type: 90-DAY

Route: Gavage

Species/Strain: Rat/F 344/N

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Test Compound: Methylene blue trihydrate

CAS Number: 7220-79-3

Date Report Requested: 10/15/2014

Time Report Requested: 18:02:41

First Dose M/F: NA / NA

Lab: BAT

**F 344/N Rat Female
200 MG/KG**

| DAY ON TEST | | | | | | | | | | |
|-------------|---|---|---|---|---|---|---|---|---|---|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| ANIMAL ID | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 |
| | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 0 |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 |

***TOTALS**

| | | | | | | | | | | | | |
|----------------------------------|---|---|---|---|---|---|---|---|---|---|----|-----|
| Lymph Follic, Depletion Cellular | 2 | 1 | 2 | 2 | 2 | 1 | 1 | 1 | 1 | 2 | 9 | 1.6 |
| Thymus | + | + | + | + | + | + | + | + | + | + | 10 | |

Integumentary System

| | | | | | | | | | | | |
|---------------|---|---|---|---|---|---|---|---|---|---|----|
| Mammary Gland | + | + | + | + | + | + | + | + | + | + | 10 |
| Skin | + | + | + | + | + | + | + | + | + | + | 10 |

Musculoskeletal System

| | | | | | | | | | | | |
|------|---|---|---|---|---|---|---|---|---|---|----|
| Bone | + | + | + | + | + | + | + | + | + | + | 10 |
|------|---|---|---|---|---|---|---|---|---|---|----|

Nervous System

| | | | | | | | | | | | |
|-------|---|---|---|---|---|---|---|---|---|---|----|
| Brain | + | + | + | + | + | + | + | + | + | + | 10 |
|-------|---|---|---|---|---|---|---|---|---|---|----|

Respiratory System

| | | | | | | | | | | | |
|----------------------------|---|---|---|---|---|---|---|---|---|---|-------|
| Lung | + | + | + | + | + | + | + | + | + | + | 10 |
| Hemorrhage, Acute, Diffuse | 3 | | | | | | | | | | 1 3.0 |
| Nose | + | + | + | + | + | + | + | + | + | + | 10 |
| Inflammation, Chronic | | | | | | | | 1 | | | 1 1.0 |
| Trachea | + | + | + | + | + | + | + | + | + | + | 10 |

Special Senses System

NONE

Urinary System

| | | | | | | | | | | | |
|--------|---|---|---|---|---|---|---|---|---|---|----|
| Kidney | + | + | + | + | + | + | + | + | + | + | 10 |
|--------|---|---|---|---|---|---|---|---|---|---|----|

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M ..Missing tissue

X ..Lesion present

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Species/Strain: Rat/F 344/N

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Test Compound: Methylene blue trihydrate

CAS Number: 7220-79-3

Date Report Requested: 10/15/2014

Time Report Requested: 18:02:41

First Dose M/F: NA / NA

Lab: BAT

F 344/N Rat Female
200 MG/KG

| DAY ON TEST | | | | | | | | | | |
|-------------|---|---|---|---|---|---|---|---|---|---|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| ANIMAL ID | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 1 | 9 | 2 | 9 | 9 | 7 | 2 | 9 | 9 | 9 |
| | 4 | 2 | 7 | 2 | 2 | 1 | 7 | 2 | 2 | 2 |

*TOTALS

Urinary Bladder

+ + + + + + + + + +

10

** END OF REPORT **

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ ..Tissue examined microscopically

M ..Missing tissue

X ..Lesion present

A ..Autolysis precludes evaluation

I ..Insufficient tissue

BLANK ..Not examined microscopically

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1) Minimal 3) Moderate

2) Mild 4) Marked